

## WHAT IS CLAIMED IS:

37. A therapeutic method for preventing hypertension, said method comprising administering to a subject susceptible to such disorder a therapeutically-effective amount of an active compound of Formula I:

$$R_{1} \xrightarrow[R_{9}]{R_{11}} \xrightarrow[R_{2}]{R_{12}} \xrightarrow[N]{R_{3}} \xrightarrow[N]{R_{4}} \xrightarrow[N]{R_{5}} \xrightarrow[N]{R_{6}} \xrightarrow[N]{OH} \xrightarrow[R_{7}]{OH} \xrightarrow[N]{R_{12}} \xrightarrow[N]{R_{12}} \xrightarrow[N]{R_{12}} \xrightarrow[N]{R_{12}} \xrightarrow[N]{R_{12}} \xrightarrow[N]{R_{12}} \xrightarrow[N]{R_{13}} \xrightarrow[N]{R_{14}} \xrightarrow[N]{R_{15}} \xrightarrow[N]{N} \xrightarrow[N} \xrightarrow[N]{N} \xrightarrow[N]{N} \xrightarrow[N} \xrightarrow[N]{N} \xrightarrow[N]{N} \xrightarrow[N} \xrightarrow[N]{N}$$

wherein A is selected from methylene, CO, SO and SO2; wherein X is selected from oxygen atom, methylene and  $\sim NR_{10}$  with  $R_{10}$ selected from hydrido, alkyl and benzyl; wherein each of R1 and R9 is a group independently selected from hydrido, alkyl, cycloalkyl, alkoxyacyl, haloalkyl, alkoxycarbonyl, benzyloxycarbonyl, loweralkanoyl, haloalkylacyl, phenyl, benzyl, naphthyl, and naphthylmethyl, any one of which groups having a substitutable position may be optionally substituted with one or more radicals selected from alkyl, alkoxy, alkenyl, alkynyl, halo, haloalkyl, cyano and phenyl, and wherein the nitrogen atom to which R<sub>1</sub> and R<sub>9</sub> are attached may be combined with oxygen to form an N-oxide; wherein R2 is selected from hydrido, alkyl, dialkylaminoalkyl, alkylacylaminoalkyl, benzyl and cycloalkyl; wherein R3 is selected from alkyl, cycloalkylalkyl, acylaminoalkyl, phenylalkyl, naphthylmethyl, aryl, heterocyclicalkyl and heterocycliccycloalkyl, wherein the cyclic portion of any of said phenylalkyl, naphthylmethyl, aryl, heterocyclicalkyl and heterocycliccycloalkyl groups may be substituted by one or more radicals selected from halo, hydroxy, alkoxy and alkyl; wherein each of R4 and R6 is independently selected from hydrido, alkyl, benzyl and cycloalkyl; wherein each of R5 and R8 is independently selected from

$$-(CH_2)_{q} \begin{bmatrix} R_{13} \\ C \\ R_{14} \end{bmatrix}_{p} C \equiv C$$

wherein V is selected from hydrido, alkyl, cycloalkyl, haloalkyl, benzyl and phenyl; wherein each of R13 and R14 is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, cycloalkyl, phenyl, heterocyclic, heterocyclicalkyl and heterocycliccycloalkyl; wherein R7 is selected from substituted or unsubstituted alkyl, cycloalkyl, phenyl, cycloalkylalkyl and phenylalkyl, any one of which may be substituted with one or more groups selected from alkyl, hydroxy, alkoxy, halo, haloalkyl, alkenyl, alkynyl and cyano; wherein each of R11 and R12 is independently selected from hydrido, alkyl, haloalkyl, dialkylamino and phenyl; and wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

38. The method of Claim 37 wherein A is selected from methylene, CO, SO and SO2; wherein X is selected from oxygen atom, methylene and  $NR_{10}$  with  $R_{10}$  selected from hydrido, alkyl and benzyl; wherein each of  $R_1$  and  $R_9$  is independently selected from hydrido, lower alkyl, haloalkyl, cycloalkyl, alkoxycarbonyl, benzyloxycarbonyl, loweralkanoyl, alkoxyacyl, phenyl and benzyl, and wherein the nitrogen atom to which  $R_1$  and  $R_9$  are attached may be combined with oxygen to form an N-oxide; wherein each of  $R_2$ ,  $R_4$  and  $R_6$  is independently selected from hydrido and alkyl; wherein  $R_3$  is selected from phenylalkyl, naphthylmethyl, cyclohexylalkyl, cyclopentylalkyl, heteroarylalkyl and heteroarylcycloalkyl; wherein each of  $R_5$  and  $R_8$  is independently selected from

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$$--\left(CH_{2}\right)_{q}\begin{bmatrix} R_{13} \\ C \\ R_{14} \\ P \end{bmatrix}_{p} C \equiv C-V$$

wherein V is selected from hydrido, alkyl, haloalkyl, benzyl and phenyl; wherein each of R<sub>13</sub> and R<sub>14</sub> is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, cycloalkyl, heteroaryl, heteroarylalkyl and heteroarylcycloalkyl; wherein R<sub>7</sub> is selected from substituted or unsubstituted cyclohexylmethyl and benzyl, either one of which may be substituted with one or more groups selected from alkyl, hydroxy, alkoxy, halo and haloalkyl; wherein each of R<sub>11</sub> and R<sub>12</sub> is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

methylene, CO, SO and SO2; wherein X is selected from oxygen atom, methylene and NR<sub>10</sub> with R<sub>10</sub> selected from hyrido, alkyl and benzyl; wherein each of R<sub>1</sub> and R<sub>9</sub> is independently selected from hydrido, alkyl, alkoxyacyl, haloalkyl, alkoxycarbonyl, benzyloxycarbonyl and benzyl, and wherein the nitrogen atom to which R<sub>1</sub> and R<sub>9</sub> are attached may be combined with oxygen to form an N-oxide; wherein each of R<sub>2</sub>, R<sub>4</sub> and R<sub>6</sub> is independently selected from hydrido and alkyl; wherein R<sub>3</sub> is selected from benzyl, phenethyl, cyclohexylmethyl, phenpropyl, pyrrolidinyl, piperidinyl, pyrrolidinylmethyl, piperidinylmethyl, pyrazolemethyl, pyrazoleethyl, pyridylmethyl, pyridylethyl, thiazoleethyl, imidazolemethyl, imidazoleethyl, thienylmethyl, furanylmethyl, furanylethyl,

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oxazolemethyl, oxazoleethyl, isoxazolemethyl, isoxazoleethyl, pyridazinemethyl, pyridazineethyl, pyrazinemethyl and pyrazineethyl; wherein each of R5 and R8 is independently selected from

$$-\left(CH_{2}\right)_{q} \begin{bmatrix} R_{13} \\ C \\ R_{14} \end{bmatrix}_{p} C \equiv C-V$$

wherein V is selected from hydrido, alkyl and haloalkyl; wherein each of  $R_{13}$  and  $R_{14}$  is a radical independently selected from hydrido, alkyl, alkenyl, alkynyl, thiazole and thiazolemethyl; wherein  $R_{7}$  is cyclohexylmethyl; wherein each of  $R_{11}$  and  $R_{12}$  is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero or one; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

40. The method of Claim 39 wherein A is selected from CO and SO2; wherein X is selected from oxygen atom, methylene and  $NR_{10}$  with  $R_{10}$  selected from hydrido and methyl; wherein each of  $R_{1}$  and  $R_{9}$  is independently selected from hydrido, lower alkyl, alkoxyacyl, alkoxycarbonyl, benzyloxycarbonyl, haloalkyl and benzyl, and wherein the nitrogen atom to which  $R_{1}$  and  $R_{9}$  are attached may be combined with oxygen to form an N-oxide; wherein  $R_{2}$  is selected from hydrido, methyl, ethyl and isopropyl; wherein  $R_{3}$  is selected from benzyl, phenethyl, cyclohexylmethyl, pyrrolidinyl, piperidinyl, pyrrolidinyl, piperidinylmethyl, thiazolemethyl, pyrazolemethyl, pyridylmethyl, pyridylmethyl, thiazolemethyl, thiazoleethyl, imidazolemethyl, imidazoleethyl, thienylmethyl, furanylmethyl, furanylethyl, oxazolemethyl, isoxazolemethyl, isoxazoleethyl, isoxazoleethyl, isoxazoleethyl,

pyridazinemethyl, pyridazineethyl, pyrazinemethyl and pyrazineethyl; wherein each of R4 and R6 is independently selected from hydrido and methyl; wherein each of R5 and R8 is independently selected from

$$-(CH_2)_{q} \begin{bmatrix} R_{13} \\ C \\ R_{14} \end{bmatrix}_{p} C \equiv C-V$$

wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R<sub>13</sub> and R<sub>14</sub> is a radical independently selected from hydrido, alkyl and alkynyl; wherein R<sub>7</sub> is cyclohexylmethyl; wherein each of R<sub>11</sub> and R<sub>12</sub> is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

41. The method of Claim 40 wherein A is selected from CO and SO2; wherein X is selected from oxygen atom and methylene; wherein each of R<sub>1</sub> and R<sub>9</sub> is independently selected from hydrido, methyl, ethyl, n-propyl, isopropyl, benzyl, b, b, b-trifluoroethyl, t-butyloxycarbonyl and methoxymethylcarbonyl, and wherein the nitrogen atom to which R<sub>1</sub> and R<sub>9</sub> are attached may be combined with oxygen to form an N-oxide; wherein R<sub>2</sub> is selected from hydrido, methyl, ethyl and isopropyl; wherein R<sub>3</sub> is selected from benzyl, cyclohexylmethyl, phenethyl, pyrazolemethyl, pyrazolemethyl, pyridylethyl, thiazolemethyl, thiazolemethyl, imidazolemethyl, imidazoleethyl, thienylmethyl, thienylmethyl, thienylmethyl, furanylmethyl, oxazolemethyl, pyridazinemethyl, oxazoleethyl, isoxazolemethyl, isoxazoleethyl, pyridazinemethyl,

pyridazineethyl, pyrazinemethyl and pyrazineethyl; wherein each of R5 and R8 is independently selected from

$$-(CH_2)_{q} \begin{bmatrix} R_{13} \\ C \\ R_{14} \end{bmatrix}_{p} C \equiv C-V$$

wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R<sub>13</sub> and R<sub>14</sub> is a radical independently selected from hydrido, methyl, ethyl, propyl and ethynyl; wherein R<sub>7</sub> is cyclohexylmethyl; wherein each of R<sub>4</sub> and R<sub>6</sub> is independently selected from hydrido and methyl; wherein each of R<sub>11</sub> and R<sub>12</sub> is independently selected from hydrido, alkyl, dialkylamino and phenyl; wherein m is zero; wherein n is a number selected from zero through five; wherein p is a number selected from zero through five; and wherein q is a number selected from zero through five; or a pharmaceutically-acceptable salt thereof.

42. The method of Claim 41 wherein A is selected from CO and SO2; wherein X is selected from oxygen atom and methylene; wherein each of R<sub>1</sub> and R<sub>9</sub> is a group independently selected from hydrido, methyl, ethyl, n-propyl, isopropyl, benzyl, b, b, b-trifluoroethyl, t-butyloxycarbonyl and methoxymethylcarbonyl, and wherein the nitrogen atom to which R<sub>1</sub> and R<sub>9</sub> are attached may be combined with oxygen to form an N-oxide; wherein R<sub>2</sub> is selected from hydrido, methyl, ethyl and isopropyl; wherein R<sub>3</sub> is selected from benzyl, cyclohexylmethyl, phenethyl, imidazolemethyl, pyridylmethyl and 2-pyridylethyl; wherein each of R<sub>5</sub> and R<sub>8</sub> is independently selected from

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$$-\left(CH_{2}\right)_{q} \begin{bmatrix} R_{13} \\ C \\ R_{14} \end{bmatrix}_{p} C \equiv C-V$$

wherein V is selected from hydrido, alkyl and trifluoromethyl; wherein each of R<sub>13</sub> and R<sub>14</sub> is a radical independently selected from hydrido, methyl and ethynyl; wherein R<sub>7</sub> is cyclohexylmethyl; wherein each of R<sub>4</sub> and R<sub>6</sub> is independently selected from hydrido and methyl; wherein each of R<sub>11</sub> and R<sub>12</sub> is independently selected from hydrido, alkyl and phenyl; wherein m is zero; wherein n is a number selected from zero through three; wherein p is a number selected from one through three; and wherein q is zero or one; or a pharmaceutically-acceptable salt thereof.

43. The method of Claim 42 wherein said compound is selected from compounds, their tautomers, and the pharmaceutically-acceptable esters and salts thereof, of the group consisting of

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- 44. The method of Claim 42 wherein said compound is N1-[1R\*-[[[1S,1R\*-(cyclohexylmethyl)-2S\*,3R\*-dihydroxy-hexynyl]amino]carbonyl]-3-butynyl]-N4-[2-(dimethylamino)ethyl]-N4-methyl-2S\*-(phenylmethyl)butanediamide or a pharmaceutically-acceptable salt thereof.
- 45. The method of Claim 42 wherein said compound is [1R\*-[[[1R\*-[[[1S,1R\*-(cyclohexylmethyl)-2S\*,3R\*-dihydroxy-hexynyl]amino]carbonyl]-3-butynyl]amino]carbonyl]-2-phenylethyl)[2-(dimethylamino)ethyl]methylcarbamate or a pharmaceutically-acceptable salt thereof.
  - **46**. The method of Claim 42 wherein said compound is H

or a pharmaceutically-acceptable salt thereof.

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47. The method of Claim 42 wherein said compound is

or a pharmaceutically-acceptable salt thereof.